news lggin Welcome to MESSENGER (APS Text) at USPTO ERRORS IN THE 5-APR-94 ISSUE ARE BEING CORRECTED NOTICE AND CLEAN PATENTS FOR THIS ISSUE WILL BE AVAILABLE SHORTLY NOTICE: THE UPDATE FOR USPAT 3-MAY-94 ISSUE IS NOW COMPLETE* The USPTO production files are current through: 10 Jan 1995 for U.S. Patent Text Data. 10 Jan 1995 for U.S. Current Classification data. 10 Jan 1995 for U.S. Patent Image Data. * PLEASE USE 305-9000 FOR NEW TELEPHONE NUMBER * DISCLAIMER: Neither the United States Government, nor any agency thereof, nor any of their contractors, subcontractors or employees make any warranty, expressed or implied, × including any warranty of marketability of fitness for a * particular purpose; nor assumes any legal liability or responsibility for any party's use, or the results of such, of the data. Help Desk --> 703-305-9000 The Help Desk is staffed for APS support 7 days/week. Monday through Friday: 6:30am - 9:00pm Saturday, Sunday, Holidays: 8:30am - 5:00 pm The Help Desk staff at this number will handle all APS related questions. >>>>>>>> NEW SUNDAY HOURS !!! <<<<<<<<<<< The APS is available: 6:30am - 9:00pm Monday through Friday 7:30am - 5:00pm Saturday, Sunday, Holidays APS is unavailable Thanksgiving Day, Christmas Day, and New Year's Day. => d his (FILE 'USPAT' ENTERED AT 14:43:14 ON 10 JAN 95) SET PAG SCR

138186 S PROMOT?

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40 S L2 OR L3

40 S POLYMERASE(W) III

L1

L2 L3

L4

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L6
              12 S ARNA
L7
             103 S ANTISENSE(W)RNA
L8
              55 S ANTI (W) SENSE (W) RNA
             114 S L6 OR L7
L9
             146 S L6 OR L7 OR L8
L10
L11
             114 S L1 AND L10
               6 S L4 AND L11
L12
            1167 S U6
L13
             276 S U(W)6
L14
               @ S 7SK
L15
L16
              21 S 7(W) SK
               Ø S HIRNA
L17
               1 S H1(W)RNA
L18
            2481 S U3
L19
            689 S U(W) 3
L20
             139 S MRP
L21
            1454 S L13 OR L14 OR L15 OR L16
L22
            2482 S L17 OR L18 OR L19
L23
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             827 S L20 OR L21
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              33 S L1 AND L4
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             103 S ANTISENSE (W) RNA
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             114 S L6 OR L7
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           2482 S L17 OR L18 OR L19
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             827 S L20 OR L21
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L26
             298 S L1 AND L25
               2 S L10 AND L26
L27
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L28
               1 S L12 AND L27
L29
               7 S L12 OR L27
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''DISPLAYED' IS NOT A RECOGNIZED COMMAND
=> 'Displayed L27 1-2 kwic w
''DISPLAYED' IS NOT A RECOGNIZED COMMAND
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=> d 112 2,3 cit fd rel
   5,354,854, Oct. 11, 1994, Expression system for use n plants to suppress
foreign expression and method; June E. Bourque, et al., 536/23.1, 24.1 [IMAGE
AVAILABLEJ
                                                 L12: 2 of 6
US PAT NO:
             5,354,854 [IMAGE AVAILABLE]
DATE FILED:
             Nov. 7, 1991
   5,324,643, Jun. 28, 1994, Method of conferring resistance to retroviral
infection; Wilson Greatbatch, et al., 435/91.32, 91.1, 91.3, 172.3, 240.1,
240.2; 536/23.1; 935/3, 6, 34, 70 [IMAGE AVAILABLE]
             5,324,643 [IMAGE AVAILABLE]
                                                 L12: 3 of 6
US PAT NO:
             Jul. 29, 1991
DATE FILED:
             Continuation-in-part of Ser. No. 156, 188, Feb. 16, 1988,
REL-US-DATA:
               abandoned.
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FILE 'JPOABS' ENTERED AT 15:05:38 ON 10 JAN 95
   JAPANESE PATENT
                                   ABSTRACTS
 * CURRENTLY, DATA IS LOADED THROUGH THE ABSTRACT PUBLICATION
 * DATE OF DECEMBER 31, 1993
 * THE LATEST GROUPS RECEIVED ARE: C1141 E1473, M1526 & P1652. *
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L3
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           40 S L2 OR L3
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           33 S L1 AND L4
L6
           12 S ARNA
L7
          103 S ANTISENSE (W) RNA
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L8
             55 S ANTI(W)SENSE(W)RNA
            114 S L6 OR L7
L9
L10
            146 S L6 OR L7 OR L8
            114 S L1 AND L10
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L12
              6 S L4 AND L11
           1167 S U6
L13
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L14
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L17
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              1 S H1 (W) RNA
L19
           2481 S U3
L20
            689 S U(W)3
L21
            139 S MRP
L22
           1454 S L13 OR L14 OR L15 OR L16
L23
           2482 S L17 OR L18 OR L19
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L25
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L26
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L27
              2 S L10 AND L26
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L28
              1 S L12 AND L27
L29
              7 S L12 OR L27
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             2 S POLYMERASE(W) III
             2 S L30 OR L31
L32
L33
             @ S ARNA
L34
             9 S ANTISENSE
             8 S ANTI(W) SENSE
L35
             16 S L33 OR L34 OR L35
L36
             Ø S L32 AND L36
L37
L38
             19 S U6
             5 S U(W)6
L39
             @ S 7SK
L40
             Ø S 7(W)SK
L41
             Ø S HIRNA
L42
L43
             0 S H1(W)RNA
            123 S U3
L44
            18 S U(W)3
L45
            27 S MRP
L46
            185 S L38 OR L39 OR L44 OR L45 OR L46
L47
          21278 S PROMOT?
L48
L49
              3 S L47 AND L48
              Ø S L32 AND L49
L50
=> lgo y
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'LGO' IS NOT A RECOGNIZED COMMAND

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FILE 'CA' ENTERED AT 15:23:59 ON 10 JAN 95 L1 129 S (POL(W)III)/BI,AB L2 1496 S (POLYMERASE(W) III) / BI, AB L3 1529 S L1 OR L2 L4 19 S ARNA/BI, AB L5 87 S (ANTI(W)SENSE(W)RNA)/BI,AB 1067 S (ANTISENSE(W)RNA)/BI,AB L6 1126 S L4 OR L5 OR L6 L7 9 S L3 AND L7 L8

= d 18 1-9 bib ab

ANSWER 1 OF 9

AN 121:271382 CA
TI Reduction in replication of the human immunodeficiency virus type 1 in human T cell lines by ***polymerase*** ***III*** -driven transcription of chimeric tRNA- ***antisense*** ***RNA*** genes
AU Junker, Uwe; Rittner, Karola; Homann, Matthias; Bevec, Dorian;

CA COPYRIGHT 1995 ACS

genes Junker, Uwe; Rittner, Karola; Homann, Matthias; Bevec, Dorian; Bohnlein, Ernst; Sczakiel, Georg Wien, Austria

SO Antisense Res. Dev. (1994), 4(3), 165-72 CODEN: AREDEI; ISSN: 1050-5261

DT Journal LA English

> Inhibition of human immunodeficiency virus type 1 (HIV-1) replication was demonstrated by using tat- and rev-directed antisense oligoribonucleotides 68 and 69 nucleotides in length. this study, human T-lymphoid cells were transduced with a murine amphotropic retroviral vector contg. a ***polymerase*** ***III*** -driven chimeric gene consisting of the human tRNAimet sequence and the short tat- and rev-directed antisense sequences that had been shown before to inhibit HIV-1 replication. transduced, G418-resistant human T-lymphoid Jurkat or CEM cells showed reduced replication of HIV-1 in the presence of antisense-contg. chimeric transcripts, but not with sense sequence-contq. transcripts. These results demonstrate that short ***RNA*** transcripts can be ***antisense*** inhibitory stably expressed endogenously using ***polymerase*** ***III*** promoters, which can reduce replication of HIV-1. The approach described in this work combines the advantages of short and, usually, synthetic oligonucleotides with the stable intracellular expression of inhibitory genes for HIV-1 in target cells. Considering the small size of the described chimeric ***polymerase*** ***III*** genes, it appears feasible to combine multiple antiviral genes with the currently available retroviral vectors as gene delivery systems.

L8 ANSWER 2 OF 9 CA COPYRIGHT 1995 ACS

AN 120:2271 CA

SO

IN Doglio, Alain; Lefebvre, Jean Claude; Cagnon, Laurence

PA University de Nice, Fr.

Fr. Demande, 29 pp.

AΙ FR 92-1608 920213 DT Patent LA French DNA vectors encoding anti-pathogen ***antisense*** AB or ribozyme are described. The DNA sequence encoding the anti-pathogen RNA is inserted between or adjacent to boxes A and B of a viral gene promoter. The vector can be used in treatment of microbial or viral infections. Vectors contg. DNA encoding ***RNA*** to HIV tat or rev nucleic acids ***antisense*** inserted between boxes A and B of the adenovirus VIA gene were prepd. HIV-1 replication was inhibited in CEM or MOLT-4 cells transfected with these vectors. ANSWER 3 OF 9 CA COPYRIGHT 1995 ACS L8 117:84331 CA AN Suppression of gene expression in plant cells utilizing antisense ΤI sequences transcribed by RNA ***polymerase*** ***III*** AU Bourque, June E.; Folk, William R. #2 Dep. Biochem., Univ. Missouri, Columbia, MO, 65211, USA CS SO Plant Mol. Biol. (1992), 19(4), 641-7 CODEN: PMBIDB: ISSN: 0167-4412 DT Journal LA English Inverted sequences of the chloramphenical acetyltransferase (CAT) AB reporter gene were fused to a soybean tRNAmeti gene lacking a terminator such that the tRNAmeti sequences caused the co-transcription of CAT antisense sequences by RNA ***III*** . When electroporated into carrot ***polymerase*** protoplasts, these antisense DNA constructs suppressed CAT enzyme activity expressed from co-electroporated DNAs contg. the CAT gene downstream of the cauliflower mosaic virus (CaMV) 35S RNA promoter. The most effective construct, an antisense sequence complementary to the 3' portion of the CAT gene, inhibited CAT activity five-fold greater than an antisense construct expressed by RNA polymerase II These results from the cauliflower mosaic virus 35S RNA promoter. indicate that antisense sequences transcribed by RNA ***III*** should efficiently suppress gene ***polymerase*** expression in plants. L8 ANSWER 4 OF 9 CA COPYRIGHT 1995 ACS AN 117:2159 CA Inhibition of adenovirus replication by the E1A antisense transcript TI initiated from hsp70 and VA-1 promoters Miroshnichenko, O. J.; Borisenko, A. S.; Ponomareva, T. I.; AU Tikhonenko, T. I. CS Inst. Agric. Biotechnol., Moscow, 127253, Russia #7 SO Biomed. Sci. (London) (1990), 1(3), 267-73 CODEN: BSCHE4; ISSN: 0955-9701 DT Journal LA English AB The E1A region of the adenoviral genome, important for initiation of virus infection and activation of other viral genes, was chosen as a target for engineering ***antisense*** ***RNA*** inhibit adenovirus 5 (Ad5) replication in COS-1 cell culture in vitro. The hsp70 promoter, taken from the appropriate heat-shock-protein gene of Drosophila melanogaster, and the VA-1 RNA promoter, derived from the Ad5 gene coding for low-mol.-mass VA-1 RNA and recognized by RNA ***polymerase*** ***III*** used as regulatory elements of transcription. The two types of recombinant constructs contained E1A fragments of 710 bp (hsp70 constructs) or 380 or 740 bp (VA-1 RNA constructs) in reverse orientation relative to the promoter position, as all as a transcription termination signal, the SV40 ori, and the gene

controlling Geneticin (antibiotic G418) resistance (G418R). After

FR 2687411 A1

930820

PI

of stable G418R cell lines were raised which expressed engineered asRNAs. Plating of Adjustment of known titer in monolayers of transfected COS-1 cells clearly showed strong inhibition of adenovirus replication by asRNAs: 75% with the hps70 promoter and 90% with the VA-1 RNA promoter.

#4

L8 ANSWER 5 OF 9 CA COPYRIGHT 1995 ACS

AN 114:242082 CA

TI Genetic construct for inhibiting RNA function

IN Beug, Hartmut; Birnstiel, Max L.; Cotten, Matthew; Wagner, Ernst; Kandolf, Harald

PA Boehringer Ingelheim International G.m.b.H., Fed. Rep. Ger.

SO Eur. Pat. Appl., 36 pp.

CODEN: EPXXDW

PI EP 387775 A1 900919

DS R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE

AI EP 90-104701 900313

PRAI AT 89-609 890316

DT Patent

LA German

A genetic construct for inhibiting RNA function comprises a AB ***III*** transcription unit contg. DNA which ***polymerase*** encodes an RNA-inhibiting RNA. A plasmid contg. the gene for methionine initiator tRNA of Xenopus was constructed. Into the ApaI site between the A and B boxes of the regulatory region, DNA encoding a ribozyme flanked by RNA complementary to U7 snRNA or erbB mRNA was inserted. This plasmid was introduced into chicken erythroblasts complexed to a polylysine-transferrin conjugate. the transformants, the target RNA was cleaved. The efficiency of cleavage was not affected by incorporation of the ribozyme into the tRNA structure, and the ribozyme was stabilized by its inclusion in the tRNA mol. Similar activity and stability were obtained when the ribozyme encoding sequence was incorporated into the intron of a tRNA gene.

L8 ANSWER 6 OF 9 CA COPYRIGHT 1995 ACS

AN 114:222638 CA

TI The rodent B2 sequence can affect expression when present in the transcribed region of a reporter gene

AU Bladon, Trevor S.; McBurney, Michael W.

CS Dep. Med., Univ. Ottawa, Ottawa, ON, K1H 8M5, Can.

SD Gene (1991), 98(2), 259-63

CODEN: GENED6; ISSN: 0378-1119

DT Journal

LA English

AB

The mouse B2 element is a moderately repetitive nt sequence of 180 bp transcribed by RNA ***polymerase*** ***III*** ***III***) at high levels in embryonic and ***Pol*** transformed cells. The B2 sequence is present in either orientation within the noncoding regions of a no. of genes transcribed by RNA polymerase II (Pol II). To det. if the small B2 transcripts generated by ***Pol*** ***!!!*** are natural ***RNA*** mols. which might hybridize to ***antisense*** complementary sequences present within Pol II transcripts, chimaeric reporter genes encoding Escherichia coli apt were constructed contg. a B2 repeat in either orientation within the 5'- or 3'-untranslated regions. These constructs were transfected into embryonal carcinoma (EC) cells and expression of the reporter gene was analyzed in EC cells and retinoic acid-treated EC cells, which contain high and low levels of small B2 RNAs, resp. Although the B2 sequences affected expression of the reporter gene, these effects did not appear to be due to hybridization of the small B2 RNA to the reporter transcripts. The presence of B2 sequences near a Pol II-transcribed gene can alter expression of that gene in a position- and orientation-dependent manner, suggesting these repetitive elements

may be cistacting regulators of gene expression.

- L8 ANSWER 7 OF 9 CA COPYRIGHT 1995 ACS
- AN 114:75955 CA
- TI Expression of chimeric tRNA-driven antisense transcripts renders NIH 3T3 cells highly resistant to Moloney murine leukemia virus replication
- AU Sullenger, Bruce A.; Lee, Thomas C.; Smith, Clayton A.; Ungers, Grace E.; Gilboa, Eli

#5

- CS Bone Marrow Transplant Serv., Mem. Sloan-Kettering Cancer Cent., New York, NY, 10021, USA
- SO Mol. Cell. Biol. (1990), 10(12), 6512-23 CODEN: MCEBD4; ISSN: 0270-7306
- DT Journal
- LA English
- NIH 3T3 cells infected with Moloney murine leukemia virus (MoMLV) AB express high levels of virus-specific RNA. To inhibit replication of the virus, chimeric tRNA genes encoding antisense templates were stably introduced into NIH 3T3 cells via a retroviral vector. Efficient expression of hybrid tRNA-MoMLV antisense transcripts and inhibition of MoMLV replication were dependent on the use of a particular type of retroviral vector, the double-copy vector, in which the chimeric tRNA gene was inserted in the 3' long terminal MoMLV replication was inhibited up to 97% in cells ***antisense*** ***RNA*** corresponding to the expressing gag gene and less than 2-fold in cells expressing ***antisense*** ***RNA*** corresponding to the pol gene. RNA and protein analyses suggest that inhibition was exerted at the level of translation. These results suggest that RNA ***polymerase*** ***!!!*** -based antisense inhibition systems can be used to inhibit highly expressed viral genes and render cells resistant to viral replication via intracellular immunization strategies.

ekd da 1/11/85

- L8 ANSWER 8 OF 9 CA COPYRIGHT 1995 ACS
- AN 113:92458 CA
- TI Silkmoth chorion ***antisense*** ***RNA*** . Structural characterization, developmental regulation and evolutionary conservation
- AU Skeiky, Yasir A. W.; Iatrou, Kostas
- CS Fac. Med., Univ. Calgary, Calgary, AB, T2N 4N1, Can.
- SO J. Mol. Biol. (1990), 213(1), 53-66 CODEN: JMOBAK; ISSN: 0022-2836
- DT Journal
- LA English

AB

Choriogenic follicular cells of the silkmoth Bombyx mori contain significant quantities of ***antisense*** ***RNA*** transcribed from chorion genes. ***Antisense*** derived from a chorion gene with a high content of cysteine, HcB.12, was characterized in detail. The antisense transcripts are initiated downstream from the 3' end of HcB.12 mRNA and extend over 75% of the length of the gene, comprising its entire second exon and part of its intervening sequence. The ***antisense*** is devoid of any significant open reading frames and is not polyadenylated. These features, combined with the presence of specific sequence motifs within its transcribed and upstream region, ***RNA*** may be transcribed by suggest that ***antisense*** ***polymerase*** ***III*** . Chorion ***antisense*** is detectable only in choriogenic follicular cells and ***RNA*** appears to be coordinately regulated with chorion mRNA. Its cytoplasmic accumulation during choriogenesis parallels that of the corresponding mRNA. Although chorion mRNA is at least 5 times more ***RNA*** , the latter is ***antisense*** abundant than present as a single-stranded entity in follicular cytoplasm but can form perfect duplexes th its mRNA complement up annealing in vitro. The possible involvement of ***antisense** ***RNA transcription in the pathway that controls the programmed expression

Us Chiuriun genes en one level of offensel ipologis initiated post-transcriptional processing is discussed. ANSWER 9 OF 9 CA COP LIGHT 1995 ACS LB 107:212659 CA AN TI Inhibition of SV40 replicon function by engineered ***antisense*** ***III*** ***RNA*** transcribed by RNA ***polymerase*** AU Jennings, P. A.; Molloy, P. L. Div. Mol. Biol., CSIRO, North Ryde, 2113, Australia CS EMBO J. (1987), 6(10), 3043-7 SO CODEN: EMJODG; ISSN: 0261-4189 DT Journal LA English AB Promoters recognized by RNA ***polymerase*** ***!!!*** used to direct synthesis of RNAs of opposite polarity to the 5' end of the mRNA for the large T-antigen of SV40. A construct was made utilizing the adenovirus (human type II) VA1 gene promoter linked to 163 bp of SV40 DNA sequences cloned in antisense orientation relative to the promoter. The SV40 sequence corresponds to the 5' end of the large T-antigen gene. In addn. to the antisense constructs, control plasmids were utilized which either lacked both promoter and SV40 elements, lacked RNA ***polymerase*** ***III*** promoter elements but contained SV40 sequences, or contained the VA1 gene promoter fused to SV40 sequences in the sense The function of the various gene fusions was demonstrated in an in vitro transcription system and in vivo by S1 nuclease 5' end mapping following transfection into COS1 cells. Cotransfection of COS1 cells with the antisense gene and a plasmid contq. an SV40 origin of replication resulted in a substantial transient inhibition of SV40-replicon function when compared to control detns. (50% to nearly complete inhibition of large T-antigen dependent DNA replication for 18-36 h). These results show that an ***antisense*** ***RNA*** generated by RNA ***polymerase*** can effectively block expression of a chromosomally located gene. => d his (FILE 'HOME' ENTERED AT 15:23:51 ON 10 JAN 95) FILE 'CA' ENTERED AT 15:23:59 ON 10 JAN 95 129 S (POL(W)III)/BI,AB L1 1496 S (POLYMERASE(W)III)/BI,AB LB L3 1529 S L1 OR L2 L4 19 S ARNA/BI,AB L5 87 S (ANTI(W) SENSE(W) RNA) /BI, AB

ekd dr

1067 S (ANTISENSE(W)RNA)/BI,AB L6 1126 S L4 OR L5 OR L6 L7 9 S L3 AND L7 L8 L9 1014 S U6/BI, AB 384 S (U(W)6)/BI,AB L10 35 S 75K/BI, AB L11 L12 6 S (7(W)SK)/BI,AB L13 Ø S H1RNA/BI, AB L14 20 S (H1(W)RNA)/BI,AB 1181 S U3/BI, AB L15 661 S (U(W)3)/BI,AB L16 320 S MRP L17 L18 320 S MRP/BI, AB L19 1373 S L9 OR L10 OR L11 OR L12 OR L13 L20 2126 S L14 OR L15 OR L16 OR L17 OR L18 L21 3432 S L19 OR L20 146539 S PROMOT?/BI,AB L22 L23 243 S L21 AND L22

1 S L7 AND L23

L24

L24 ANSWER 1 OF 1 CA COPYRIGHT 1995 ACS

AN 121:292778 CA

TI Expression constructs containing HIV inhibiting antisense sequences and their delivery by traditional means or using retrovirus expression vectors

IN Pyati, Jaqdeesh

PA Ortho Pharmaceutical Corp., USA

SO Eur. Pat. Appl., 33 pp.

CODEN: EPXXDW

PI EP 612844 A2 940831

DS R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE

AI EP 94-301315 940224

PRAI US 93-21936 930225

US 94-190350 940201

DT Patent

LA English

AB Antisense nucleotides that inhibit replication of human immunodeficiency virus are described for use in the treatment and prophylaxis of AIDS. The constructs are administered to the patient by traditional pharm. methods, or through the use of recombinant retrovirus delivery systems. The retrovirus delivery systems may be target-specific. Such targeting is accomplished by modifying the envelope of the retrovirus to contain sequences for which a receptor or ligand exists on the target. The construction of a no. of antisense expression vectors is demonstrated. Two of these vectors were packaged using an amphotropic cell line and the virus used to infect a T-lymphoblastoid cell line. The cells were shown to transcribe the antisense message and were 75-80% resistant to challenge with HIV-1.

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CA SUBSCRIBER PRICE	-4.20	-4.2Ø

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